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NOTICE OF ALLOWANCE AND FEE(S) DUE

LIFE TECHNOLOGIES CORPORATION C/O INTELLEVATE P.O. BOX 52050 MINNEAPOLIS, MN 55402 EXAMINER
SISSON, BRADLEY L

ART UNIT PAPER NUMBER
1634

DATE MAILED: 05/03/2012

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
09/613,903	07/11/2000	Heather J. Jordan	IVGN 187.1 CON	1446

TITLE OF INVENTION: NUCLEIC ACID LADDERS

	APPLN. TYPE	SMALL ENTITY	ISSUE FEE DUE	PUBLICATION FEE DUE	PREV. PAID ISSUE FEE	TOTAL FEE(S) DUE	DATE DUE
Ī	nonprovisional	NO	\$1740	\$0	\$0	\$1740	08/03/2012

THE APPLICATION IDENTIFIED ABOVE HAS BEEN EXAMINED AND IS ALLOWED FOR ISSUANCE AS A PATENT. PROSECUTION ON THE MERITS IS CLOSED. THIS NOTICE OF ALLOWANCE IS NOT A GRANT OF PATENT RIGHTS. THIS APPLICATION IS SUBJECT TO WITHDRAWAL FROM ISSUE AT THE INITIATIVE OF THE OFFICE OR UPON PETITION BY THE APPLICANT. SEE 37 CFR 1.313 AND MPEP 1308.

THE ISSUE FEE AND PUBLICATION FEE (IF REQUIRED) MUST BE PAID WITHIN <u>THREE MONTHS</u> FROM THE MAILING DATE OF THIS NOTICE OR THIS APPLICATION SHALL BE REGARDED AS ABANDONED. <u>THIS STATUTORY PERIOD CANNOT BE EXTENDED.</u> SEE 35 U.S.C. 151. THE ISSUE FEE DUE INDICATED ABOVE DOES NOT REFLECT A CREDIT FOR ANY PREVIOUSLY PAID ISSUE FEE IN THIS APPLICATION. IF AN ISSUE FEE HAS PREVIOUSLY BEEN PAID IN THIS APPLICATION (AS SHOWN ABOVE), THE RETURN OF PART B OF THIS FORM WILL BE CONSIDERED A REQUEST TO REAPPLY THE PREVIOUSLY PAID ISSUE FEE TOWARD THE ISSUE FEE NOW DUE.

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B. If the status above is to be removed, check box 5b on Part B - Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and twice the amount of the ISSUE FEE shown above, or

If the SMALL ENTITY is shown as NO:

A. Pay TOTAL FEE(S) DUE shown above, or

B. If applicant claimed SMALL ENTITY status before, or is now claiming SMALL ENTITY status, check box 5a on Part B - Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and 1/2 the ISSUE FEE shown above.

II. PART B - FEE(S) TRANSMITTAL, or its equivalent, must be completed and returned to the United States Patent and Trademark Office (USPTO) with your ISSUE FEE and PUBLICATION FEE (if required). If you are charging the fee(s) to your deposit account, section "4b" of Part B - Fee(s) Transmittal should be completed and an extra copy of the form should be submitted. If an equivalent of Part B is filed, a request to reapply a previously paid issue fee must be clearly made, and delays in processing may occur due to the difficulty in recognizing the paper as an equivalent of Part B.

III. All communications regarding this application must give the application number. Please direct all communications prior to issuance to Mail Stop ISSUE FEE unless advised to the contrary.

IMPORTANT REMINDER: Utility patents issuing on applications filed on or after Dec. 12, 1980 may require payment of maintenance fees. It is patentee's responsibility to ensure timely payment of maintenance fees when due.

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Complete and send this form, together with applicable fee(s), to: Mail Mail Stop ISSUE FEE

Commissioner for Patents P.O. Box 1450

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maintenance fee notifications. Note: A certificate of mailing can only be used for domestic mailings of the Fee(s) Transmittal. This certificate cannot be used for any other accompanying papers. Each additional paper, such as an assignment or formal drawing, must CURRENT CORRESPONDENCE ADDRESS (Note: Use Block 1 for any change of address) have its own certificate of mailing or transmission. 05/03/2012 Certificate of Mailing or Transmission LIFE TECHNOLOGIES CORPORATION I hereby certify that this Fee(s) Transmittal is being deposited with the United States Postal Service with sufficient postage for first class mail in an envelope addressed to the Mail Stop ISSUE FEE address above, or being facsimile transmitted to the USPTO (571) 273-2885, on the date indicated below. C/O INTELLEVATE P.O. BOX 52050 MINNEAPOLIS, MN 55402 (Depositor's name (Signature (Date APPLICATION NO. FILING DATE FIRST NAMED INVENTOR ATTORNEY DOCKET NO. CONFIRMATION NO. IVGN 187.1 CON 09/613,903 07/11/2000 Heather J. Jordan 1446 TITLE OF INVENTION: NUCLEIC ACID LADDERS PUBLICATION FEE DUE DATE DUE ISSUE FEE DUE PREV. PAID ISSUE FEE TOTAL FEE(S) DUE APPLN, TYPE SMALL ENTITY NO \$1740 \$0 \$0 \$1740 08/03/2012 nonprovisional **EXAMINER** ART UNIT CLASS-SUBCLASS SISSON, BRADLEY L 536-023100 1634 1. Change of correspondence address or indication of "Fee Address" (37 CFR 1.363). 2. For printing on the patent front page, list (1) the names of up to 3 registered patent attorneys or agents OR, alternatively, ☐ Change of correspondence address (or Change of Correspondence Address form PTO/SB/122) attached. (2) the name of a single firm (having as a member a registered attorney or agent) and the names of up to "Fee Address" indication (or "Fee Address" Indication form PTO/SB/47; Rev 03-02 or more recent) attached. Use of a Customer Number is required. 2 registered patent attorneys or agents. If no name is listed, no name will be printed. 3. ASSIGNEE NAME AND RESIDENCE DATA TO BE PRINTED ON THE PATENT (print or type) PLEASE NOTE: Unless an assignee is identified below, no assignee data will appear on the patent. If an assignee is identified below, the document has been filed for recordation as set forth in 37 CFR 3.11. Completion of this form is NOT a substitute for filing an assignment. (B) RESIDENCE: (CITY and STATE OR COUNTRY) (A) NAME OF ASSIGNEE 4b. Payment of Fee(s): (Please first reapply any previously paid issue fee shown above) 4a. The following fee(s) are submitted: lssue Fee A check is enclosed. ☐ Publication Fee (No small entity discount permitted) Payment by credit card. Form PTO-2038 is attached. The Director is hereby authorized to charge the required fee(s), any deficiency, or credit any overpayment, to Deposit Account Number ______ (enclose an extra copy of this fo Advance Order - # of Copies _ (enclose an extra copy of this form). 5. Change in Entity Status (from status indicated above) ☐ b. Applicant is no longer claiming SMALL ENTITY status. See 37 CFR 1.27(g)(2). a. Applicant claims SMALL ENTITY status. See 37 CFR 1.27. NOTE: The Issue Fee and Publication Fee (if required) will not be accepted from anyone other than the applicant; a registered attorney or agent; or the assignee or other party in interest as shown by the records of the United States Patent and Trademark Office. Authorized Signature Date Typed or printed name Registration No. This collection of information is required by 37 CFR 1.311. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to take 12 minutes to complete, including gathering, preparing, and an apparation. Community is governed by 53 0.3.C. 122 and 57 CFR 1.14. Inis collection is estimated to take 12 minutes to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, Virginia 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, Virginia 22313-1450.

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09/613,903	07/11/2000	Heather J. Jordan	IVGN 187.1 CON	1446
65482 75	90 05/03/2012	EXAMINER		
LIFE TECHNOLOGIES CORPORATION C/O INTELLEVATE P.O. BOX 52050 MINNEAPOLIS, MN 55402			SISSON, BRADLEY L	
			ART UNIT	PAPER NUMBER
			1634	

DATE MAILED: 05/03/2012

Determination of Patent Term Adjustment under 35 U.S.C. 154 (b)

(application filed on or after May 29, 2000)

The Patent Term Adjustment to date is 0 day(s). If the issue fee is paid on the date that is three months after the mailing date of this notice and the patent issues on the Tuesday before the date that is 28 weeks (six and a half months) after the mailing date of this notice, the Patent Term Adjustment will be 0 day(s).

If a Continued Prosecution Application (CPA) was filed in the above-identified application, the filing date that determines Patent Term Adjustment is the filing date of the most recent CPA.

Applicant will be able to obtain more detailed information by accessing the Patent Application Information Retrieval (PAIR) WEB site (http://pair.uspto.gov).

Any questions regarding the Patent Term Extension or Adjustment determination should be directed to the Office of Patent Legal Administration at (571)-272-7702. Questions relating to issue and publication fee payments should be directed to the Customer Service Center of the Office of Patent Publication at 1-(888)-786-0101 or (571)-272-4200.

Privacy Act Statement

The Privacy Act of 1974 (P.L. 93-579) requires that you be given certain information in connection with your submission of the attached form related to a patent application or patent. Accordingly, pursuant to the requirements of the Act, please be advised that: (1) the general authority for the collection of this information is 35 U.S.C. 2(b)(2); (2) furnishing of the information solicited is voluntary; and (3) the principal purpose for which the information is used by the U.S. Patent and Trademark Office is to process and/or examine your submission related to a patent application or patent. If you do not furnish the requested information, the U.S. Patent and Trademark Office may not be able to process and/or examine your submission, which may result in termination of proceedings or abandonment of the application or expiration of the patent.

The information provided by you in this form will be subject to the following routine uses:

- 1. The information on this form will be treated confidentially to the extent allowed under the Freedom of Information Act (5 U.S.C. 552) and the Privacy Act (5 U.S.C 552a). Records from this system of records may be disclosed to the Department of Justice to determine whether disclosure of these records is required by the Freedom of Information Act.
- 2. A record from this system of records may be disclosed, as a routine use, in the course of presenting evidence to a court, magistrate, or administrative tribunal, including disclosures to opposing counsel in the course of settlement negotiations.
- 3. A record in this system of records may be disclosed, as a routine use, to a Member of Congress submitting a request involving an individual, to whom the record pertains, when the individual has requested assistance from the Member with respect to the subject matter of the record.
- 4. A record in this system of records may be disclosed, as a routine use, to a contractor of the Agency having need for the information in order to perform a contract. Recipients of information shall be required to comply with the requirements of the Privacy Act of 1974, as amended, pursuant to 5 U.S.C. 552a(m).
- 5. A record related to an International Application filed under the Patent Cooperation Treaty in this system of records may be disclosed, as a routine use, to the International Bureau of the World Intellectual Property Organization, pursuant to the Patent Cooperation Treaty.
- 6. A record in this system of records may be disclosed, as a routine use, to another federal agency for purposes of National Security review (35 U.S.C. 181) and for review pursuant to the Atomic Energy Act (42 U.S.C. 218(c)).
- 7. A record from this system of records may be disclosed, as a routine use, to the Administrator, General Services, or his/her designee, during an inspection of records conducted by GSA as part of that agency's responsibility to recommend improvements in records management practices and programs, under authority of 44 U.S.C. 2904 and 2906. Such disclosure shall be made in accordance with the GSA regulations governing inspection of records for this purpose, and any other relevant (i.e., GSA or Commerce) directive. Such disclosure shall not be used to make determinations about individuals.
- 8. A record from this system of records may be disclosed, as a routine use, to the public after either publication of the application pursuant to 35 U.S.C. 122(b) or issuance of a patent pursuant to 35 U.S.C. 151. Further, a record may be disclosed, subject to the limitations of 37 CFR 1.14, as a routine use, to the public if the record was filed in an application which became abandoned or in which the proceedings were terminated and which application is referenced by either a published application, an application open to public inspection or an issued patent.
- 9. A record from this system of records may be disclosed, as a routine use, to a Federal, State, or local law enforcement agency, if the USPTO becomes aware of a violation or potential violation of law or regulation.

	Application No.	Applicant(s)			
	09/613,903	JORDAN, HEATHE	·B .I		
Notice of Allowability	Examiner	Art Unit			
	Bradley L. Sisson	1634			
The MAILING DATE of this communication appe All claims being allowable, PROSECUTION ON THE MERITS IS herewith (or previously mailed), a Notice of Allowance (PTOL-85) NOTICE OF ALLOWABILITY IS NOT A GRANT OF PATENT RI of the Office or upon petition by the applicant. See 37 CFR 1.313	(OR REMAINS) CLOSED in the or other appropriate communing GHTS. This application is subtand MPEP 1308.	nis application. If not includication will be mailed in due pject to withdrawal from issu	led course. THIS ue at the initiative		
1. A This communication is responsive to Appeal Brief of 02/10/2	* *				
 An election was made by the applicant in response to a rest the restriction requirement and election have been incorporate 		uring the interview on	_;		
3. 🛮 The allowed claim(s) is/are <u>141,150,155-159 and 162-173</u> .					
 4. ☐ Acknowledgment is made of a claim for foreign priority unde a) ☐ All b) ☐ Some* c) ☐ None of the: 					
1. Certified copies of the priority documents have		NI.			
2. Certified copies of the priority documents have			ation from the		
 Copies of the certified copies of the priority doc International Bureau (PCT Rule 17.2(a)). 	cuments have been received in	n this national stage applica	ation from the		
* Certified copies not received:					
Applicant has THREE MONTHS FROM THE "MAILING DATE" noted below. Failure to timely comply will result in ABANDONM THIS THREE-MONTH PERIOD IS NOT EXTENDABLE.		reply complying with the re	quirements		
 A SUBSTITUTE OATH OR DECLARATION must be submit INFORMAL PATENT APPLICATION (PTO-152) which give 			OTICE OF		
6. CORRECTED DRAWINGS (as "replacement sheets") must	t be submitted.				
(a) \square including changes required by the Notice of Draftspers	on's Patent Drawing Review (PTO-948) attached			
1) 🗌 hereto or 2) 🔲 to Paper No./Mail Date					
(b) ☐ including changes required by the attached Examiner's Paper No./Mail Date					
Identifying indicia such as the application number (see 37 CFR 1. each sheet. Replacement sheet(s) should be labeled as such in the			e back) of		
 DEPOSIT OF and/or INFORMATION about the deposit of B attached Examiner's comment regarding REQUIREMENT FC 					
Attachment(s) 1. ☑ Notice of References Cited (PTO-892)		rmal Patent Application			
2. Notice of Draftperson's Patent Drawing Review (PTO-948)	6. ⊠ Interview Sum Paper No./M:				
3. Information Disclosure Statements (PTO/SB/08), Paper No./Mail Date		Paper No./Mail Date <u>04/30/2012</u> . 7. ☑ Examiner's Amendment/Comment			
 Examiner's Comment Regarding Requirement for Deposit of Biological Material 	8. ⊠ Examiner's St 9. □ Other	atement of Reasons for All	owance		
/Bradley L. Sisson/ Primary Examiner, Art Unit 1634					

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EXAMINER'S AMENDMENT

1. An examiner's amendment to the record appears below. Should the changes and/or additions be unacceptable to applicant, an amendment may be filed as provided by 37 CFR 1.312. To ensure consideration of such an amendment, it MUST be submitted no later than the payment of the issue fee.

- 2. Authorization for this examiner's amendment was given in a telephone interview with Jonathan Aumais, Ph. D., Reg. No. 65,710, on 30 April 2012.
- 3. The application has been amended as follows:
 - 141. (Currently amended): A nucleic acid ladder consisting essentially of a plurality of double stranded nucleic acid fragments, each fragment having a size in base pairs of between 20 kb and 100 base pairs, a copy number, a mass, and a relative mass wherein the mass of each fragment is the size in base pairs of the fragment multiplied by the copy number of the fragment, wherein the relative mass of each fragment is the mass of the fragment divided by the sum of the masses of all of the fragments, wherein the relative mass of any one fragment of the plurality is no more than 3 time the relative mass of any other fragment of the plurality, wherein at least two of the plurality of nucleic acid fragments have a size greater than 1 kb, and wherein at least two of the plurality of nucleic acid fragments have a size less than 1 kb, wherein the nucleic acid ladder lacks a highlight fragment.
 - 165. (Currently amended): A nucleic acid ladder comprising a plurality of double stranded nucleic acid <u>fragmentsmolecules</u>, wherein three or more of the <u>fragmentsmolecules</u> are of a size selected from the group consisting of:
 - (a) 100 base pairs,
 - (b) 200 base pairs,
 - (c) 300 base pairs,
 - (d) 400 base pairs,
 - (e) 500 base pairs,
 - (f) 650 base pairs,
 - (g) 850 base pairs, and

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(h) 1650 base pairs; and

wherein two or more of the fragmentsmolecules are of a size selected from the group consisting of:

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- (a) 1 kilobase pairs,
- (b) 2 kilobase pairs,
- (c) 3 kilobase pairs,
- (d) 4 kilobase pairs, and
- (e) 5 kilobase pairs;

wherein each fragment has a copy number, a mass, and a relative mass wherein the mass of each fragment is the size in base pairs of the fragment multiplied by the copy number of the fragment, wherein the relative mass of each fragment is the mass of the fragment divided by the sum of the masses of all of the fragments, wherein the relative mass of any one fragment of the plurality is no more than 3 time the relative mass of any other fragment of the plurality—a copy number of each of the molecules is such that each molecule has a relative mass that is no more than three times the relative mass of another molecule.

- 169. (Currently amended): A nucleic acid ladder comprising a plurality of double stranded nucleic acid molecules fragments, wherein three or more of the molecules fragments are of a size selected from the group consisting of:
 - (a) 100 base pairs,
 - (b) 200 base pairs,
 - (c) 300 base pairs,
 - (d) 400 base pairs,
 - (e) 500 base pairs,
 - (f) 650 base pairs,
 - (g) 850 base pairs, and
 - (h) 1650 base pairs; and

wherein two or more of the <u>molecules fragments</u> are of a size selected from the group consisting of:

- (a) 1 kilobase pairs,
- (b) 2 kilobase pairs,
- (c) 3 kilobase pairs,

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(d) 4 kilobase pairs, and

(e) 5 kilobase pairs;

wherein a copy number of the molecules is such that each molecule has a relative mass that is no more than three times the relative mass of another molecule each fragment has a copy number, a mass, and a relative mass wherein the mass of each fragment is the size in base pairs of the fragment multiplied by the copy number of the fragment, wherein the relative mass of each fragment is the mass of the fragment divided by the sum of the masses of all of the fragments, wherein the relative mass of any one fragment of the plurality is no more than 3 time the relative mass of any other fragment of the plurality and one or both of the following:

i)—wherein the nucleic acid ladder further comprises at least one highlight fragment having a size in the range of 100 base pairs to 5 kilobase pairs and having a relative mass that is three times greater than the relative mass of other molecules in the composition, or;

ii) wherein at least one of the three or more molecules is a highlight fragment having a relative mass that is three times greater than the relative mass of the other molecules in the composition.

- 4. The following is an examiner's statement of reasons for allowance: Claims 141, 150, 155-159, and 162-173 are allowable over the teaching of US Patent 7,132,520 B2 (Hartley), and Life Technologies, "GIBCO BRL Product Catalogue and Reference Guide 1995-1996," the closest prior art of record.
- 5. Hartley, while published on 07 November 2006, claims benefit of priority to application filed 28 October 1993, and thusly qualifies as 102(e)-type art.
- 6. Hartley claims a "nucleic acid marker ladder," kit comprising same, and related method of making. The nucleic acid marker ladder is an endonuclease digestion. The ladder of Hartley, while requiring anywhere from 3 to 10 fragments, "wherein the size of each said fragment in base pairs is a multiple of an integer between 10 and 100," does not require that the fragments

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have the same "relative mass," as is required of the presently-claimed nucleic acid ladder.

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Accordingly, Hartley does not teach or reasonably suggest the claimed invention.

- Life Technologies advertises for sale a 100 bp DNA ladder at page 14-3 which comprises a plurality of bands ranging from 100 bp to 2 Kb. However, the bands at 600 and 2 Kb are arguably a "highlight fragment," which is expressly excluded from claim 141, and claims 150, 155-159, and 162-164 which depend therefrom. Further, the 100 bp DNA ladder does not have two fragments in the range of 1 kb to 5 kb where said fragments are not a highlight fragment (requirement of independent claims 165 an 169). Rather, the prior art teaches a plurality of fragments where there are two fragments in the range of 1 kb and 2 kb, but one of the fragments is a highlight fragment. Accordingly, the prior art does not ach or fairly suggest the claimed combination of DNA fragments of which the claimed nucleic acid ladder is comprised.
- 8. Any comments considered necessary by applicant must be submitted no later than the payment of the issue fee and, to avoid processing delays, should preferably accompany the issue fee. Such submissions should be clearly labeled "Comments on Statement of Reasons for Allowance."
- 9. Any inquiry concerning this communication or earlier communications from the examiner should be directed to Bradley L. Sisson whose telephone number is (571)272-0751. The examiner can normally be reached on 6:30 a.m. to 5 p.m., Monday through Thursday.
- 10. If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Dave T. Nguyen can be reached on (571) 272-0731. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

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11. Information regarding the status of an application may be obtained from the Patent

Application Information Retrieval (PAIR) system. Status information for published applications

may be obtained from either Private PAIR or Public PAIR. Status information for unpublished

applications is available through Private PAIR only. For more information about the PAIR

system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR

system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would

like assistance from a USPTO Customer Service Representative or access to the automated

information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

/Bradley L. Sisson/

Primary Examiner, Art Unit 1634